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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/901,484	07/09/2001	Daniel Cohen	GEN-T111XC3D2	6608
23557	23557 7590 07/15/2005		EXAMINER	
SALIWANCHIK LLOYD & SALIWANCHIK A PROFESSIONAL ASSOCIATION		FREDMAN, JEFFREY NORMAN		
PO BOX 1429			ART UNIT	PAPER NUMBER
	LE, FL 32614-2950		1637	

DATE MAILED: 07/15/2005

Please find below and/or attached an Office communication concerning this application or proceeding.

	من				
	Application No.	Applicant(s)			
Office Action Occurrence	09/901,484	COHEN ET AL.			
Office Action Summary	Examiner	Art Unit			
	Jeffrey Fredman	1637			
The MAILING DATE of this communication app Period for Reply	oears on the cover sheet with the c	orrespondence address			
A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) FROM THE MAILING DATE OF THIS COMMUNICATION. - Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication. - If the period for reply specified above is less than thirty (30) days, a reply within the statutory minimum of thirty (30) days will be considered timely. - If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication. - Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). - Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b). Status					
1) Responsive to communication(s) filed on <u>05 J</u>	uly 2005.				
2a) ☐ This action is FINAL . 2b) ☐ This	action is non-final.				
3) Since this application is in condition for allowated closed in accordance with the practice under the condition of the					
Disposition of Claims					
4) Claim(s) 50-53,56-60,63,64,67,68 and 71-84 i	s/are pending in the application.				
4a) Of the above claim(s) is/are withdra	wn from consideration.				
5) Claim(s) is/are allowed.					
6) Claim(s) 50-53,56-59,63,64,67,68,71,72 and 7	74-84 is/are rejected.	·			
7)⊠ Claim(s) 60 and 73 is/are objected to.					
8) Claim(s) are subject to restriction and/o	or election requirement.				
Application Papers					
9)☐ The specification is objected to by the Examine	•	•			
10)☐ The drawing(s) filed on is/are: a)☐ acc					
Applicant may not request that any objection to the					
Replacement drawing sheet(s) including the correct					
11) The oath or declaration is objected to by the E	xaminer. Note the attached Office	Action or form PTO-152.			
Priority under 35 U.S.C. §§ 119 and 120		•			
12) Acknowledgment is made of a claim for foreignal All by Some * c) None of: 1. Certified copies of the priority document 2. Certified copies of the priority document 3. Copies of the certified copies of the priority application from the International Burea * See the attached detailed Office action for a list 13) Acknowledgment is made of a claim for domest	ts have been received. Its have been received in Applicationity documents have been receive In (PCT Rule 17.2(a)). In of the certified copies not receive	ion No ed in this National Stage			
since a specific reference was included in the fir 37 CFR 1.78. a) The translation of the foreign language product of a claim for demost	ovisional application has been rec	ceived.			
14) Acknowledgment is made of a claim for domestic priority under 35 U.S.C. §§ 120 and/or 121 since a specific reference was included in the first sentence of the specification or in an Application Data Sheet. 37 CFR 1.78.					
Attachment(s)					
1) Notice of References Cited (PTO-892) 2) Notice of Draftsperson's Patent Drawing Review (PTO-948) 3) Information Disclosure Statement(s) (PTO-1449) Paper No(s)	5) 🔲 Notice of Informal F	(PTO-413) Paper No(s) Patent Application (PTO-152)			

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DETAILED ACTION

Priority

1. The current application claims priority to a series of cases dating back to 1997. However, the claims are not given priority to applications 08/996,306 and 60/099,658 because in the current application SEQ ID NO: 179 is 56,520 nucleotides while in those parent applications, the largest sequences were 56,516 nucleotides. Consequently, there is no possibility that these applications provide full descriptive support for SEQ ID NO: 179, and priority to these applications is denied. Therefore, for purposes of prior art, the priority date of this application is limited to 09/218,207, filed December 22,1998, which provides the full 56,520 nucleotides of SEQ ID NO: 179.

Claim Rejections - 35 USC § 102

2. The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless -

- (a) the invention was known or used by others in this country, or patented or described in a printed publication in this or a foreign country, before the invention thereof by the applicant for a patent.
- (b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.
- 3. The rejection of claims 50, 59 and 61-66 under 35 U.S.C. 102(b) as being anticipated by Weier et al (Hum. Genet. (1991) 87:489-494) as evidenced by Genbank Accession No. AC100813 (March 2003) is withdrawn in view of the amendment..
- 4. Claims 50-52, 56-59, 63, 64, 67, 68, 71-72 and 74-84 are rejected under 35 U.S.C. 102(a) and (b) as being anticipated by Osoegawa et al (Genomics (1998)52:1-8)

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as evidenced by Genbank Accession No. AC009631 and an email from Pieter de Jong (attached).

Osoegawa teaches synthesis of BAC chromosome libraries (abstract). In particular, Osoegawa teaches the synthesis of a particular BAC library termed RPCI-11 (also called RP11) which was grown in bacterial host cells using recombinant vectors which were placed onto filters and the nucleic acid was isolated (page 2 and page 3, column 1).

A specific isolated BAC, RP11-372K15 which is in a composition of this library and which is at a specific location on the arrays sold by the BACPAC consortium, comprises 2010 contiguous nucleotides from nucleotides that overlap position 51333, 51435, 51468, 51515, 51557, 51566, 51632, 51666, 52016, 52096, 52151, 52282, 52348, 5241, 52580, 5271, 52772, 52860 and 53092 of SEQ ID NO: 179. As shown by the alignment below, this BAC has close match with the reference sequence.

```
Score = 3828 bits (1931), Expect = 0.0
Identities = 2009/2010 (99%), Gaps = 0/2010 (0%)
Strand=Plus/Plus
SEQ ID NO: 179 51233
{\tt AATATCAAAAGAGTCGGTGTGAACCTTGGTTGGACCCCAAGTTCACAAGATTTTTAAGGT}
                                                  51292
AC009631
              446
AATATCAAAAGAGTCGGTGTGAACCTTGGTTGGACCCCAAGTTCACAAGATTTTTAAGGT
                                                  505
SEQ ID NO: 179 51293
{\tt GATGAGAGCCTGCAGACATTCTGCCTAGATTTACTAGCGTGTGCCTTTTGCCTGCTTCTC}
                                                  51352
GATGAGAGCCTGCAGACATTCTGCCTAGATTTACTAGCGTGTGCCTTTTGCCTGCTTCTC 565
```

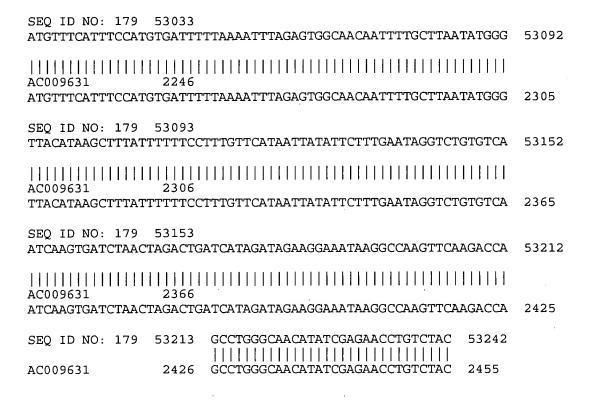
SEQ ID NO: 179 51353 TTTGATTTCACAGAATATTCATTCAGAAGTCGCGTTTCTGTAGTGTGGTGGATTCCCACT	51412
	625
SEQ ID NO: 179 51413 GGGCTCTGGTCCTTCCCTTGGATCCCGTCAGTGGTGCTCCTCAGCGGCTTGCACGTAGAC	51472
	685
SEQ ID NO: 179 51473 TTGCTAGGAAGAAATGCAGAGCCAGCCTGTGCTGCCCACTTTCAGAGTTGAACTCTTTAA	51532
	745
SEQ ID NO: 179 51533 GCCCTTGTGAGTGGGCTTCACCAGCTACTGCAGAGGCATTTTGCATTTGTCTGTC	51592
	805
SEQ ID NO: 179 51593 AAGTTCACCTTCTCAAGCCAGTGAAATACAGACTTAATTCGTCATGACTGAACGAATTTG	51652
	865
SEQ ID NO: 179 51653 TTTATTTCCCATTAGGTTTAGTGGAGCTACACATTAATATGTATCGCCTTAGAGCAAGAG	51712
	925
SEQ ID NO: 179 51713 CTGTGTTCCAGGAACCAGATCACGATTTTTAGCCATGGAACAATATATCCCATGGGAGAA	51772
	005
CTGTGTTCCAGGAACCAGATCACGATTTTTAGCCATGGAACAATATATCCCATGGGAGAA	985

SEQ ID NO: 179 51773 GACCTTTCAGTGTGAACTGTTCTATTTTTGTGTTATAATTTAAACTTCGATTTCCTCATA	51832
	1045
SEQ ID NO: 179 51833 GTCCTTTAAGTTGACATTTCTGCTTACTGCTACTGGATTTTTGCTGCAGAAATATATCAG	51892
	1105
SEQ ID NO: 179 51893 TGGCCCACATTAAACATACCAGTTGGATCATGATAAGCAAAATGAAAGAAA	51952
	1165
SEQ ID NO: 179 51953 AGGGAAAATTAAGTGACTGTTTACACTGCTTCTCCCATGCCAGAGAATAAACTCTTTCA	52012
AC009631 1166 AGGGAAAATTAAGTGACTGTTTACACTGCTTCTCCCATGCCAGAGAATAAACTCTTTCA	1225
SEQ ID NO: 179 52013 AGCATCATCTTTGAAGAGTCGTGTGGTGTGAATTGGTTTGTGTACATTAGAATGTATGCA	52072
AC009631 1226 AGC G TCATCTTTGAAGAGTCGTGTGTGTGTAATTGGTTTGTGTACATTAGAATGTATGCA	1285
SEQ ID NO: 179 52073 CACATCCATGGACACTCAGGATATAGTTGGCCTAATAATCGGGGCATGGGTAAAACTTAT	52132
	52152
CACATCCATGGACACTCAGGATATAGTTGGCCTAATAATCGGGGCATGGGTAAAACTTAT	1345
SEQ ID NO: 179 52133 GAAAATTTCCTCATGCTGAATTGTAATTTTCTCTTACCTGTAAAGTAAAATTTAGATCAA	52192
	1405
GAAAATTICCICATGCIGAATTGTAATTTTCTCTTACCTGTAAAGTAAA	1403

SEQ ID NO: 179 52193 TTCCATGTCTTTGTTAAGTACAGGGATTTAATATATTTTGAATATAATGGGTATGTTCTA	52252
	1465
SEQ ID NO: 179 52253 AATTTGAACTTTGAGAGGCAATACTGTTGGAATTATGTGGATTCTAACTCATTTTAACAA	52312
	1525
SEQ ID NO: 179 52313 GGTAGCCTGACCTGCATAAGATCACTTGAATGTTAGGTTTCATAGAACTATACTAATCTT	52372
	1585
SEQ ID NO: 179 52373 CTCACAAAAGGTCTATAAAATACAGTCGTTGAAAAAAATTTTGTATCAAAATGTTTGGAA	52432
	1645
SEQ ID NO: 179 52433 AATTAGAAGCTTCTCCTTAACCTGTATTGATACTGACTTGAATTATTTTCTAAAATTAAG	52492
	1705
SEQ ID NO: 179 52493 AGCCGTATACCTACCTGTAAGTCTTTTCACATATCATTTAAACTTTTGTTTG	52552
	1765°
SEQ ID NO: 179 52553 CTGATTTACAGCTTAGTTATTAATTTTCTTTATAAGAATGCCGTCGATGTGCATGCTTT	52612
	1825

SEQ ID NO: 179 52613	
TATGTTTTCAGAAAAGGGTGTGTTTGGATGAAAGTAAAAAAAA	52672
	1885
	1000
SEQ ID NO: 179 52673 TGTCTCTAATGGCTGTGCTGTTTAACATTTTTTGACCCTAAAATTCACCAACAGTCTCCC	52732
TGTCTCTAATGGCTGTGCTGTTTAACATTTTTTGACCCTAAAATTCACCAACAGTCTCCC	1945
SEQ ID NO: 179 52733 AGTACATAAAATAGGCTTAATGACTGGCCCTGCATTCTTCACAATATTTTTCCCTAAGCT	52792
AGTACATAAAATAGGCTTAATGACTGGCCCTGCATTCTTCACAATATTTTTCCCTAAGCT	2005
SEQ ID NO: 179 52793 TTGAGCAAAGTTTTAAAAAAATACACTAAAATAATCAAAACTGTTAAGCAGTATATTAGT	52852
TTGAGCAAAGTTTTAAAAAAATACACTAAAATAATCAAAACTGTTAAGCAGTATATTAGT	2065
SEQ ID NO: 179 52853 TTGGTTATATAAATTCATCTGCAATTTATAAGATGCATGGCCGATGTTAATTTGCTTGGC	52912
AC009631 2066 TTGGTTATATAAATTCATCTGCAATTTATAAGATGCATGGCCGATGTTAATTTGCTTGGC	2125
SEQ ID NO: 179 52913 AATTCTGTAATCATTAAGTGATCTCAGTGAAACATGTCAAATGCCTTAAATTAACTAAGT	52972
AATTCTGTAATCATTAAGTGATCTCAGTGAAACATGTCAAATGCCTTAAATTAACTAAGT	2185
SEQ ID NO: 179 52973 TGGTGAATAAAAGTGCCGATCTGGCTAACTCTTACACCATACATA	53032
TGGTGAATAAAAGTGCCGATCTGGCTAACTCTTACACCATACATA	2245

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The above alignment, meets several of the elements of claim 50. The sequence meets element (a) because there are more than 1000 consecutive nucleotides of SEQ ID NO: 179, specifically nucleotides 52017 to 53242 which comprise 1225 consecutive nucleotides of SEQ ID NO: 179. This sequence also comprises the complement claimed in (d) and meets (e) for all of positions 51333, 51435, 51468, 51515, 51557, 51566, 51632, 51666, 52016, 52096, 52151, 52282, 52348, 5241, 52580, 5271, 52772, 52860 and 53092 of SEQ ID NO: 179, where the central T is position 32703 of SEQ ID NO: 179, where N is one of these listed positions and X is within the range of 8-30, including 8, 10, 12, 15, 20 or 25.

The email of Pieter de Jong indicates that filters from the RPCI-11 library were first publicly available, used and sold on August 1, 1997.

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With regard to claim 51, Osoegawa teaches that the sequences were in Bac vectors (see page 1, column 2, subheading "BAC/PAC vector preparation").

With regard to claim 52, Osoegawa teaches that the vectors were in bacterial host cells (see page 2, column 1).

With regard to claim 56, Osoegawa teaches that the sequences were in vectors, and the vector sequence can function as a label for the detection of the target sequence (see page 1, column 2. To explain, a DNA sequence may itself be a label, and frequently is used as such, since specific DNA can be detected by hybridization).

With regard to claims 57-58, Osoegawa teaches that the oligonucleotide is attached, indirectly, to a solid support (see page 2, column 1 and email, where filters were sold).

With regard to claims 59, 63-64, Osoegawa teaches a library which would comprise the RPCI-11 library, which Genbank Accession No. AC009631 shows has 1225 contiguous nucleotides in the claimed region (see alignment above).

With regard to claims 67-68, the phrase "consisting essentially of" is treated as equivalent to "comprising" since no specific definition of the term is provided and therefore Osoegawa also anticipates these claims.

With regard to claims 71-72, 74-84, Osoegawa teaches oligonucleotides of 1225 contiguous nucleotides which comprise a contiguous span of more than 1000 nucleotides overlapping positions 52626-53599 (see alignment above).

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5. Claims 50-52, 56-59, 63, 64, 67 and 68 are rejected under 35 U.S.C. 102(a) and (b) as being anticipated by Osoegawa et al (Genomics (1998)52:1-8) as evidenced by Genbank Accession No. AC068274 and an email from Pieter de Jong (attached).

Osoegawa teaches synthesis of BAC chromosome libraries (abstract). In particular, Osoegawa teaches the synthesis of a particular BAC library termed RPCI-11 (also called RP11) which was grown in bacterial host cells using recombinant vectors which were placed onto filters and the nucleic acid was isolated (page 2 and page 3, column 1).

A specific isolated BAC, RP11-119B15 which is in a composition of this library and which is at a specific location on the arrays sold by the BACPAC consortium, comprises 17 contiguous nucleotides from nucleotides that overlap position 32703 of SEQ ID NO: 179. As shown by the alignment below, this BAC has close match with the reference sequence.

The above alignment, where the central T is position 32703 of SEQ ID NO: 179, meets claim 50, element (e), since there is a sequence comprising a contiguous span of nucleotides where N is 32703, where X is equal to 8 and where Y is equal to 8.

The email of Pieter de Jong indicates that filters from the RPCI-11 library were first publicly available, used and sold on August 1, 1997.

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With regard to claim 51, Osoegawa teaches that the sequences were in Bac vectors (see page 1, column 2, subheading "BAC/PAC vector preparation").

With regard to claim 52, Osoegawa teaches that the vectors were in bacterial host cells (see page 2, column 1).

With regard to claim 56, Osoegawa teaches that the sequences were in vectors, and the vector sequence can function as a label for the detection of the target sequence (see page 1, column 2. To explain, a DNA sequence may itself be a label, and frequently is used as such, since specific DNA can be detected by hybridization).

With regard to claims 57-58, Osoegawa teaches that the oligonucleotide is attached, indirectly, to a solid support (see page 2, column 1 and email, where filters were sold).

With regard to claims 59, 63-64, Osoegawa teaches a library which would comprise the RPCI-11 library, which Genbank Accession No. AC068274shows has 543 contiguous nucleotides in the claimed region (see alignment above).

With regard to claims 67-68, the phrase "consisting essentially of" is treated as equivalent to "comprising" since no specific definition of the term is provided and therefore Osoegawa also anticipates these claims.

Claim Rejections - 35 USC § 103

- 6. The following is a quotation of 35 U.S.C. 103(a) which forms the basis for all obviousness rejections set forth in this Office action:
 - (a) A patent may not be obtained though the invention is not identically disclosed or described as set forth in section 102 of this title, if the differences between the subject matter sought to be patented and the prior art are such that the subject matter as a whole would have been obvious at the time the invention was made to a person having ordinary skill in the art to which said subject matter pertains. Patentability shall not be negatived by the manner in which the invention was made.

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- 7. This application currently names joint inventors. In considering patentability of the claims under 35 U.S.C. 103(a), the examiner presumes that the subject matter of the various claims was commonly owned at the time any inventions covered therein were made absent any evidence to the contrary. Applicant is advised of the obligation under 37 CFR 1.56 to point out the inventor and invention dates of each claim that was not commonly owned at the time a later invention was made in order for the examiner to consider the applicability of 35 U.S.C. 103(c) and potential 35 U.S.C. 102(e), (f) or (g) prior art under 35 U.S.C. 103(a).
- 8. Claim 53 is rejected under 35 U.S.C. 103(a) as being unpatentable over Osoegawa et al (Genomics (1998)52:1-8) in view of Capecchi et al (Science (1989) 244:1288-1292).

Osoegawa teaches vectors that comprise sequences of interest as discussed above.

Capecchi teaches the use of homologous recombination to form host cells and mammals (see page 1280, figure 1, for example).

It would have been prima facie obvious to one of ordinary skill in the art at the time the invention was made to screen each of the sequences of Osoegawa for functional activity using the homologous recombination method of Capecchi since Capecchi states "Targeted disruption of these genes may not only reveal the phenotypes associated with inactivation of the individual genes, but through epistasis and molecular analyses, may also help define the developmental network controlling early mouse morphogenesis (see page 1292, column 1)." Thus, an ordinary

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practitioner, interested in identifying what phenotype is associated with the sequence of the sequences of Osoegawa would have been motivated by Capecchi to use targeted disruption in order to define the phenotype of the genes with which the sequence of Osoegawa are associated.

Allowable Subject Matter

- 9. Claims 60 and 73 are objected to as being dependent upon a rejected base claim, but would be allowable if rewritten in independent form including all of the limitations of the base claim and any intervening claims.
- 10. The following is a statement of reasons for the indication of allowable subject matter: Claim 60 is drawn to the complete SEQ ID NO: 179. No such sequence was found in the sequence search and while the evidence from the chromosome 8 hits is that there is generally 99.8% or so alignment with hundreds of contiguous basepairs, there is no evidence that Weier is inherently identical over the entire length of SEQ ID NO: 179. Therefore, the claim to the entire sequence is novel and unobvious. With regard to claim 73, no sequences with at least 40 contiguous bases of SEQ ID NO: 179 at the specified positions were found in the sequence search. Therefore, these fragments are novel and unobvious.

Response to Arguments ...

11. Applicant's arguments with respect to the claims have been considered but are most in view of the new ground(s) of rejection necessitated by the amendment.

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Conclusion

12. Applicant's amendment necessitated the new ground(s) of rejection presented in this Office action. Accordingly, **THIS ACTION IS MADE FINAL**. See MPEP § 706.07(a). Applicant is reminded of the extension of time policy as set forth in 37 CFR 1.136(a).

A shortened statutory period for reply to this final action is set to expire THREE MONTHS from the mailing date of this action. In the event a first reply is filed within TWO MONTHS of the mailing date of this final action and the advisory action is not mailed until after the end of the THREE-MONTH shortened statutory period, then the shortened statutory period will expire on the date the advisory action is mailed, and any extension fee pursuant to 37 CFR 1.136(a) will be calculated from the mailing date of the advisory action. In no event, however, will the statutory period for reply expire later than SIX MONTHS from the date of this final action.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Jeffrey Fredman whose telephone number is (571)272-0742. The examiner can normally be reached on 6:30-3:00.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Gary Benzion can be reached on (571)272-0782. The fax phone number for the organization where this application or proceeding is assigned is 703-872-9306.

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Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free).

Jeffréy Fredman Primary Examiner